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**WO 01/74301 A1**

(54) Title: DENTAL MATERIALS WITH EXTENDABLE WORK TIME, KITS, AND METHODS

(57) Abstract: The present invention provides a dental material that includes a hardenable resin system and a first initiator system capable of hardening the hardenable resin.

**DENTAL MATERIALS WITH EXTENDABLE WORK TIME,  
KITS, AND METHODS**

**Field of the Invention**

5        This invention relates to dental materials with an extendable work time. More specifically, the dental materials have a relatively fast initial set phase and a subsequent extended flexible phase working time. Preferably, the dental materials are used as provisional oral prosthetic devices such as inlays, onlays, veneers, temporary crowns, permanent crowns, and bridges.

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**Background of the Invention**

Currently available dental materials for the fabrication of provisional restoratives include polymerizable resins, which have replaced the use of metal shells and polycarbonate temporary crowns. Resin-based crowns can be made available in different shades that can be matched to the patient's dentition. Two-part resin formulations are typically used wherein the resin constituents are separate from the initiator system. The resin constituents are typically free radically polymerizable monomers (e.g., acrylates, methacrylates, and acrylamides). The initiator system typically includes reducing agents, such as tertiary amines, and oxidizing agents, such as peroxides, as accelerators or activators for free radical polymerization and control of the rate of polymerization.

15       It is desirable to control the rate of polymerization such that the dental material has a relatively rapid initial set phase time during which time a specific hardness is attained without locking into the undercuts produced during tooth preparation, followed by an extended flexible phase work time. During this flexible phase the dental material can be removed from the mouth to allow the dental practitioner time to modify, trim, and adjust the dental material for a custom fit onto the prepared tooth. Modification and adjustment of the dental material during the extended flexible phase is desirable in order to provide a more custom fit of the provisional restorative in the mouth of the patient. Additionally, heat generated by the exothermic polymerization is controlled by removal of the dental material from the intra-oral environment. This also avoids damage to oral tissues

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and consequently provides more comfort for the patient. Before final placement in the mouth and cementing in place, the restorative is completely hardened by conventional means commonly known for free radical polymerization.

A typical procedure for making a provisional dental restorative involves the following steps. Initially, an alginate impression is taken before preparing the teeth. The impression is rinsed, set aside, and wrapped in a moist paper towel. The teeth are then prepared and the correct shade of acrylic powder is selected to match the natural teeth. An acrylic liquid resin and the acrylic polymeric powder, one of which includes a reducing agent and the other of which includes an oxidizing agent, are mixed together and placed in the impression. The impression is placed aside until the composition thickens and forms a dull appearance (approximately 45-60 seconds). Meanwhile, the prepared teeth and surrounding tissue are coated with a petroleum jelly, which ensures easy removal of the acrylic temporary from the preparation and protects the teeth and tissue from irritation by the acrylic mixture. The impression with the acrylic mixture is seated in the mouth and held in place for a sufficient time to allow it to harden to a removable state. Mixing through this stage is the initial set phase time. The acrylic material is removed from the impression and gross excess acrylic is trimmed. The acrylic material is placed in and out of the mouth while the acrylic material is in a rubbery state. This is the extended flexible phase work time. The acrylic material is removed from the mouth and set aside until the acrylic is fully cured. The fit of the acrylic restorative is checked and adapted to fit, if necessary. Excess acrylic is trimmed with an acrylic bur or stone and polished to a smooth finish. The acrylic temporary is then cemented into place.

The important time during the preparation of a restorative occurs after the resin, initiator, and reducing agent are mixed together. It is important that hardening occurs initially rapidly (during the initial set phase time), but then slows to allow a dental practitioner to place, remove, and trim the restorative material (during the extended flexible phase work time) before it continues to harden to such a state as to become unworkable. There is a need for dental materials that provide reasonable initial set phase times and flexible phase work times that are more readily controllable than conventional materials.

### Summary of the Invention

The present invention provides a dental material useful for the fabrication of dental restoratives. Such a dental material includes a resin system and a first 5 initiator system. Preferably, it also includes a filler system, and more preferably, a second initiator system. Once the components are combined to initiate a reaction, the time period during which the resultant composition solidifies to a partially hardened state is referred to herein as the initial set phase time. The partially hardened material at this point is firm enough to be removed from the mouth 10 without being permanently deformed, but not so rigid that the solidified impression locks into undercuts and becomes too hard and brittle to be trimmed. The material then has an extended flexible phase work time sufficient for the dental practitioner to place, remove, and trim the dental material before it continues to harden to such a state as to become unworkable. The dental restorative can be further hardened to 15 a rigid phase extra-orally.

In one embodiment, the dental material includes a hardenable resin system and a first initiator system capable of hardening the hardenable resin. Optionally and preferably, the dental material also includes a filler system and a second initiator system. Preferably, the hardenable resin system and the first initiator 20 system are selected such that when combined the resultant composition has an initial set phase time of less than about 180 seconds at room temperature, and an extended flexible phase working time of greater than about 85 seconds at room temperature.

The first initiator system preferably includes at least one reducing agent and 25 at least one oxidizing agent. Preferably, the oxidizing agent is a peroxide compound and the reducing agent is a tertiary aromatic amine, a mercaptan, or combinations thereof. Typically and preferably, the dental material is provided in at least two parts, wherein at least one part includes at least one reducing agent and at least one other part includes at least one oxidizing agent, which together form 30 the first initiator system.

In a preferred embodiment, the dental material includes a hardenable resin system and a first initiator system capable of hardening the hardenable resin,

wherein the first initiator system includes at least one polymerizable tertiary aromatic amine. Preferably, the polymerizable tertiary aromatic amine is selected from the group of bis-N,N-[2-(2-methacryloxyethylaminocarbonyloxy)ethyl]-p-toluidine, bis-N,N-(2-methacryloxyethyl)-p-toluidine, 4-[2(2-acrylamido-2-methylpropionyloxy)ethyl]-N,N-dimethylaniline, bis-N,N-[2-(2-acrylamido-2-methylpropionyloxy)ethyl]-p-toluidine, and mixtures thereof.

In another preferred embodiment, the dental material includes a hardenable resin system and a first initiator system capable of hardening the hardenable resin, wherein the first initiator system includes at least two peroxide oxidizing agents.

In yet another preferred embodiment, the dental material includes a hardenable resin system and a first initiator system capable of hardening the hardenable resin, wherein the first initiator system includes at least two reducing agents, at least one of which is a mercaptan.

In still another preferred embodiment, the dental material is suitable for use as a provisional restorative and includes a hardenable resin system and a first initiator system capable of hardening the hardenable resin, wherein the first initiator system includes at least two tertiary aromatic amine reducing agents.

The present invention also provides a kit for preparation of a dental restorative. The kit includes: a first container including at least one reducing agent; a second container including at least one oxidizing agent; and at least a portion of a hardenable resin system in at least one of the containers; wherein the hardenable resin system, reducing agent, and oxidizing agent are selected such that when combined the resultant composition has an initial set phase time of less than about 180 seconds at room temperature, and an extended flexible phase working time of greater than about 85 seconds at room temperature.

In another embodiment, the kit includes: a first container including at least one oxidizing agent; a second container including at least one polymerizable tertiary aromatic amine reducing agent; and a hardenable resin system in at least one of the containers.

In yet another embodiment, the kit includes: a first container including at least two peroxide oxidizing agents; a second container including at least one reducing agent; and a hardenable resin system in at least one of the containers.

In still another embodiment, the kit includes: a first container including at least one oxidizing agent; a second container including at least two reducing agents, at least one of which is a mercaptan; and a hardenable resin system in at least one of the containers.

5       The present invention also provides a method for preparing a dental restorative. The method involves: making an impression of at least one tooth; filling the impression with a hardenable composition comprising a hardenable resin system and a first initiator system, wherein the hardenable resin system and the first initiator system are selected such that when combined the resultant composition has an initial set phase time of less than about 180 seconds at room temperature, and an extended flexible phase working time of greater than about 85 seconds at room temperature; placing the impression over at least one tooth to partially harden the hardenable composition; removing the impression and partially hardened composition and removing excess hardened material to form a dental restorative; 10 and allowing the dental restorative to further harden.  
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### Definitions

The dental materials of the present invention include a hardenable resin. As used herein, "harden," "hardening," and "hardenable" is descriptive of a resin that can be polymerized and/or crosslinked.  
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"Dental material" refers to a combination of components that upon reaction form a material that is suitable for use in a patient's dentition. Herein, dental material refers to the unreacted components (e.g., various combinations of hardenable resin, initiator systems, etc.) as well as the reacted components.

25       "Initial set phase time" is the time period, as measured in seconds, during which time the dental material hardens to a hardness of 20 (i.e.,  $T_{20}$ ), as determined by measuring with a Shore A Durometer (The Shore Instrument and Manufacturing Co., New York, NY), at room temperature. Typically, the initial set phase time starts with mixing the oxidizing agent, reducing agent, and resin.

30       "Extended flexible phase time" is the time period wherein the dental material that has obtained a hardness of 20 on the shore A Durometer is allowed to harden to a hardness of no greater than 80 (i.e.,  $T_{80}$ ), as determined by measuring

with a Shore A Durometer, at room temperature. The extended flexible phase time, as measured in seconds, is obtained by subtracting T<sub>20</sub> from T<sub>80</sub>.

“Restorative” refers to an oral prosthetic device, such as an inlay, onlay, a veneer, a single unit crown, a splint, a crown lining material for prefabricated crowns, an implant, or multiple unit bridges.

“Provisional Restorative” is a temporary placement intended for eventual removal and replacement by a more long term or permanent device. Typically, a provisional restorative is intended to be in the intra-oral environment for no greater than about a year, preferably for no greater than about 6 months, and more preferably no greater than about 30 days.

#### Brief Description of Drawings

Figure 1 is a chart of Shore A Hardness values of three comparative dental materials.

Figure 2 is a chart of the initial set phase and extended flexible phase times of three comparative dental materials.

#### Detailed Description of Preferred Embodiments of the Invention

This invention addresses use of initiator systems and hardenable resins to design a dental material with a controllable initial set phase. The dental material is useful as a restorative, such as inlays, onlays, veneers, crowns, and bridges, for example. Preferably, the dental material is useful as a provisional restorative, such as a temporary crown and temporary bridge.

A dental material of the present invention includes a unique combination of a resin system and a first initiator system. Preferably, it also includes a filler system and a second initiator system. By judicious selection of these components, particularly the resin system and first initiator system, the dental material upon mixing of the components preferably exhibits an initial hardening to 20 on a Shore A Durometer in no greater than about 180 seconds (i.e., an initial set phase time). This initial hardening of the composition not only captures the topography of the prepared tooth, but also the composition is still easily removable from the mouth as a flexible or rubbery solid. Preferably, for at least about 85 seconds after removing

the partially hardened dental material from the mouth the dental practitioner has time to trim the dental material to provide a custom fit onto the prepared tooth. During this time period, which is referred to herein as the extended flexible phase working time, the dental material preferably continues to harden to a hardness of 5 80 on a Shore A Durometer. After the dental material has been suitably modified to provide a custom fit in the mouth during this extended flexible phase, the dental material is allowed to harden further, preferably to a fully hardened state. Herein, the times of the initial set phase and the extended flexible working phase are measured at room temperature (approximately 20-25°C). When the hardening occurs in the mouth, these times may be slightly shorter as a result of the higher 10 temperature of the mouth (e.g., 30-38°C).

Dental materials of the type described in detail herein are normally supplied to the dental practitioner in at least two separate parts. At least one of these parts contains at least one reducing agent, which is one component of a first initiator 15 system and at least one other part contains at least one oxidizing agent, which is a second component of a first initiator system. In using the dental material, the dentist mixes the portions to produce a hardenable composition. The dentist places the freshly prepared hardenable composition into a tooth template (impression). The inventors have discovered that by carefully selecting the individual 20 components of the dental material one or more of the following problems is solved: the material hardens too quickly to an unworkable state; the material hardens too quickly and locks into tooth undercuts; the material does not harden to a workable state in a sufficient period of time; and the material produces too much heat during curing for patient comfort.

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### *Resin System*

The resin system includes one or more hardenable organic resins capable of forming a hardened material having sufficient strength and hydrolytic stability to render them suitable for use in the oral environment. A suitable organic resin is 30 hardenable (e.g., polymerizable) by a free radical mechanism and includes monomers, oligomers, and/or polymers. Preferably, at least some of the monomers, oligomers, and/or polymers include ethylenic unsaturation and are

capable of undergoing addition polymerization. A suitable resin preferably includes at least one ethylenically unsaturated monomer (i.e., includes at least one carbon-carbon double bond).

Examples of suitable resin components include: mono-, di- or poly-  
5 acrylates and methacrylates such as methyl acrylate, methyl methacrylate, ethyl acrylate, isopropyl methacrylate, n-hexyl acrylate, stearyl acrylate, allyl acrylate, glycerol diacrylate, glycerol triacrylate, ethyleneglycol diacrylate, diethyleneglycol diacrylate, triethyleneglycol dimethacrylate, 1,3-propanediol diacrylate, 1,3-  
10 propanediol dimethacrylate, trimethylolpropane triacrylate, 1,2,4-butanetriol trimethacrylate, 1,4-cyclohexanediol diacrylate, pentaerythritol triacrylate, pentaerythritol tetraacrylate, pentaerythritol tetramethacrylate, sorbitol hexacrylate, bis[1-(2-acryloxy)]-p-ethoxyphenyldimethylmethane, bis[1-(3-acryloxy-2-  
hydroxy)]-p-propoxyphenyldimethylmethane, tris(hydroxyethylisocyanurate)  
15 trimethacrylate, the bis-acrylates and bis-methacrylates of polyethylene glycols of molecular weight 200-500, copolymerizable mixtures of acrylated monomers such as those of U.S. Pat. No. 4,652,274, and acrylated oligomers such as those of U.S. Pat. No. 4,642,126; unsaturated amides such as methylene bis-acrylamide, methylene bis-methacrylamide, 1,6-hexamethylene bis-acrylamide, diethylene triamine tris-acrylamide, and beta-methacrylamidoethyl methacrylate; and vinyl  
20 compounds such as styrene, diallyl phthalate, divinyl succinate, divinyl adipate, and divinylphthalate. Mixtures of two or more monomers, oligomers, and/or polymers in the resin system can be used if desired.

#### *First Initiator System*

25 The first initiator system is a redox initiator that includes an oxidizing agent (such as a peroxide) and a reducing agent (such as an aromatic amine). Combining the redox couple results in the generation of an initiating species (such as free radicals or cations) capable of causing curing (e.g., polymerization and/or crosslinking) of a hardenable resin. Preferably, the redox couples of this invention are activated at temperatures below about 40°C. Generally, the redox couple is partitioned into separate reactive compositions prior to use and then subsequently mixed at the time of use to generate the desired initiating species. Selection of the  
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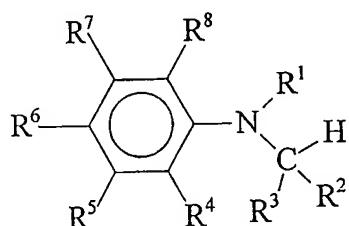
redox couple is governed by several criteria. For example, a desirable oxidizing agent is one that is sufficiently oxidizing in nature to oxidize the reducing agent, but not excessively oxidizing that it may prematurely react with resins or other components with which it may be combined during storage. Similarly, a desirable reducing agent is one that is sufficiently reducing in nature to readily react with the preferred oxidizing agent, but not excessively reducing in nature such that it may reduce the resin or other components with which it may be combined during storage. Oxidation or reduction of the resin with an inappropriate reducing agent or oxidizing agent, respectively, could result in an unstable system that would prematurely polymerize and subsequently provide a limited shelf life. Thus, suitable redox couples individually provide good shelf-life, and then, when combined together, generate the desired initiating species for hardening of the resin system.

Suitable oxidizing agents include peroxide compounds (i.e., peroxy compounds), including hydrogen peroxide as well as inorganic and organic peroxide compounds (e.g., "per" compounds or salts with peroxyanions). Examples of suitable oxidizing agents include, but are not limited to: peroxides such as benzoyl peroxide, phthaloyl peroxide, substituted benzoyl peroxides, acetyl peroxide, caproyl peroxide, lauroyl peroxide, cinnamoyl peroxide, acetyl benzoyl peroxide, methyl ethyl ketone peroxide, sodium peroxide, hydrogen peroxide, di-tert butyl peroxide, tetraline peroxide, urea peroxide, and cumene peroxide; hydroperoxides such as p-methane hydroperoxide, di-isopropyl-benzene hydroperoxide, tert- butyl hydroperoxide, methyl ethyl ketone hydroperoxide, and 1-hydroxy cyclohexyl hydroperoxide-1, ammonium persulfate, sodium perborate, sodium perchlorate, potassium persulfate, etc.; ozone, ozonides, etc. These oxidizing agents may be used alone or in admixture with one another. Benzoyl peroxide is the preferred oxidizing agent. One or more oxidizing agents may be present in the first initiator system in an amount sufficient to provide initiation of the hardening process. Preferably, this includes about 0.01 weight percent (wt-%) to about 4.0 wt-%, and more preferably about 0.05 wt-% to about 1.0 wt-%, based on the total weight of all components of the dental material.

The first initiator system also includes a reducing agent with one or more functional groups for activation of the oxidizing agent and initiation of hardening. The functional groups provide both a fast and a slow rate of activation for the initiation of hardening for the initial set phase and extended flexible phase, respectively. Preferably, the functional groups are selected from amines, mercaptans, or mixtures thereof. If more than one functional group is present, they may be part of the same compound or provided by different compounds.

A preferred reducing agent is a tertiary aromatic amine. An example of the type of useful tertiary amines to the invention is

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wherein each R group can be H or an organic group that does not adversely effect the initiation of hardening of the dental material. Preferably, the organic groups would not sterically or electronically hinder the function of the reducing agent.

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Examples of such compounds are disclosed in WO 97/35916, published October 2, 1997.

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As used herein, the term "organic group" means a hydrocarbon group (with optional elements other than carbon and hydrogen, such as oxygen, nitrogen, sulfur, and silicon) that is classified as an aliphatic group, cyclic group, or combination of aliphatic and cyclic groups (e.g., alkaryl and aralkyl groups). It also includes functional groups such as carboxylic acid groups, esters, amides, and the like. The term "aliphatic group" means a saturated or unsaturated linear or branched hydrocarbon group. This term is used to encompass alkyl, alkenyl, and alkynyl groups, for example. The term "alkyl group" means a saturated linear or branched hydrocarbon group including, for example, methyl, ethyl, isopropyl, t-butyl, heptyl, dodecyl, octadecyl, amyl, 2-ethylhexyl, and the like. The term "alkenyl group" means an unsaturated linear or branched hydrocarbon group with

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one or more carbon-carbon double bonds, such as a vinyl group. The term "alkynyl group" means an unsaturated linear or branched hydrocarbon group with one or more carbon-carbon triple bonds. The term "cyclic group" means a closed ring hydrocarbon group that is classified as an alicyclic group, aromatic group, or heterocyclic group. The term "alicyclic group" means a cyclic hydrocarbon group having properties resembling those of aliphatic groups. The term "aromatic group" or "aryl group" means a mono- or polynuclear aromatic hydrocarbon group. The term "heterocyclic group" means a closed ring hydrocarbon in which one or more of the atoms in the ring is an element other than carbon (e.g., nitrogen, oxygen, sulfur, etc.).

Substitution is anticipated on the organic groups of the compounds described herein. As a means of simplifying the discussion and recitation of certain terminology used throughout this application, the terms "group" and "moiety" are used to differentiate between chemical species that allow for substitution or that may be substituted and those that do not allow or may not be so substituted. Thus, when the term "group" is used to describe a chemical substituent, the described chemical material includes the unsubstituted group and that group with O, N, Si, or S atoms, for example, in the chain (as in an alkoxy group) as well as carbonyl groups or other conventional substitution. Where the term "moiety" is used to describe a chemical compound or substituent, only an unsubstituted chemical material is intended to be included. For example, the phrase "alkyl group" is intended to include not only pure open chain saturated hydrocarbon alkyl substituents, such as methyl, ethyl, propyl, t-butyl, and the like, but also alkyl substituents bearing further substituents known in the art, such as hydroxy, alkoxy, alkylsulfonyl, halogen atoms, cyano, nitro, amino, carboxyl, etc. Thus, "alkyl group" includes ether groups, haloalkyls, nitroalkyls, carboxyalkyls, hydroxyalkyls, sulfoalkyls, etc. On the other hand, the phrase "alkyl moiety" is limited to the inclusion of only pure open chain saturated hydrocarbon alkyl substituents, such as methyl, ethyl, propyl, t-butyl, and the like.

Referring to the tertiary amine described above, preferably R<sup>1</sup> is an aliphatic group and R<sup>2</sup> and R<sup>3</sup> are independently (i.e., they may be the same or different) H, aromatic and/or aliphatic groups (preferably including up to 20 carbon atoms).

Preferably, only one of R<sup>2</sup> and R<sup>3</sup> is an aromatic group. More preferably, R<sup>1</sup> is an alkyl group (preferably including up to 10 carbon atoms) optionally substituted with hydroxy groups, and R<sup>2</sup> and R<sup>3</sup> are H or alkyl groups (preferably including up to 10 carbon atoms) optionally substituted with hydroxyl groups. For certain preferred embodiments, R<sup>1</sup>, R<sup>2</sup>, and R<sup>3</sup> can also include a polymerizable functional group that will react with the functional groups of the resin. Preferably, at least one of R<sup>1</sup>, R<sup>2</sup>, and R<sup>3</sup> includes a functional group such as an acrylate, methacrylate, acrylamide, vinyl, or other functional group present in the resins described above.

Preferably, R<sup>4</sup>, R<sup>5</sup>, R<sup>6</sup>, R<sup>7</sup>, and R<sup>8</sup> are independently H or aliphatic groups (preferably including up to 20 carbon atoms). More preferably, R<sup>4</sup>, R<sup>5</sup>, R<sup>6</sup>, R<sup>7</sup>, and R<sup>8</sup> are independently H or alkyl groups (preferably including up to 10 carbon atoms) optionally substituted with hydroxy groups. For certain preferred embodiments, R<sup>4</sup>, R<sup>5</sup>, R<sup>6</sup>, R<sup>7</sup>, and R<sup>8</sup> can also include a polymerizable functional group that will react with the functional groups of the resin. Preferably, at least one of R<sup>4</sup>, R<sup>5</sup>, R<sup>6</sup>, R<sup>7</sup>, and R<sup>8</sup> includes a functional group such as an acrylate, methacrylate, acrylamide, vinyl, or other functional group present in the resins described above.

Particularly preferred aromatic tertiary amines are N,N-bis(2-hydroxyethyl)-p-toluidine (DHEPT), 2-(4-dimethylaminophenyl)ethyl alcohol (DMAPE), 4-tert butyl dimethyl aniline. Other compounds that are suitable include compounds derived from DMAPE with di- or multi-functional acid compounds such as adipic acid, sebacic acid, 1,3,5-benzene tricarboxylic acid, 1,2,4,5-benzene tetracarboxylic acid, and the like, or DMAPE with di-or multi-functional isocyanates such as hexamethylene diisocyanate, isophorone diisocyanate, and desmodur N-330 (a trifunctional isocyanate).

The tertiary amines may be polymerizable. Particularly preferred polymerizable aromatic tertiary amines include, but are not limited to, an adduct of IEM (2-isocyanatoethylmethacrylate) with N,N-bis(2-hydroxyethyl)-p-toluidine (DHEPT-di-IEM or bis-N,N-[2-(2-methacryloxyethylaminocarbonyloxy)ethyl]-p-toluidine), an adduct of DMAPE with VDM (2-vinyl-4,4-dimethylazlactone) (DMAPE-VDM or 4-[2(2-acrylamido-2-methylpropionyloxy)ethyl]-N,N-dimethylaniline), an adduct of a methacrylate di-ester with DHEPT (DHEPT-di-

ester or bis-N,N-(2-methacryloyloxyethyl)-p-toluidine), and an adduct of DHEPT with VDM (DHEPT-di-VDM or bis-N,N-[2-(2-acrylamido-2-methylpropionyloxy)ethyl]-p-toluidine).

Another preferred reducing agent is a mercaptan, which can include aromatic and/or aliphatic groups, and optionally polymerizable groups. Preferred mercaptans have a molecular weight greater than about 200 as these mercaptans have less odor. Particularly preferred mercaptans are iso-octylthioglycoate (IOTG) and pentaerythritol tetrakis (3-mercaptopropionate) (PETMP).

The tertiary amines and mercaptans may be used alone or in admixture with one another. For example, the first initiator system can include one tertiary aromatic amine and one mercaptan, two tertiary aromatic amines, two mercaptans, one polymerizable tertiary aromatic amine, and the like. Other reducing agents, such as sulfinic acids, formic acid, ascorbic acid, hydrazines, and salts thereof, can also be used herein to initiate free radical polymerization. Preferably, however, the first initiator system includes a tertiary aromatic amine, a mercaptan, or a mixture thereof. Such reducing agents can function as both a component of the first initiator system and a component of the second initiator system.

If two or more reducing agents are used, they are preferably chosen such that at least one has a faster rate of activation than the other(s). That is, one causes a faster rate of initiation of the hardening of the resin than the other(s). Significantly, this provides for the two hardening stages referred to herein as the initial set phase and the extended flexible phase.

Electrochemical oxidation potentials of reducing agents and reduction potentials of oxidizing agents are useful tools for predicting the effectiveness of a suitable redox couple. For example, the reduction potential of the oxidant (i.e., oxidizing agent) benzoyl peroxide is approximately -0.16 volts vs. a saturated calomel electrode (SCE). Similarly, the oxidation potential (vs. SCE) for a series of amines has been previously established as follows: dihydroxyethyl-p-toluidine ((DHEPT), 0.76 volt), 4-t-butyl dimethylaniline ((t-BDMA), 0.77 volt), 4-dimethylaminophenethanol ((DMAPE), 0.78 volt), triethylamine ((TEA, 0.88 volt), 3-dimethylaminobenzoic acid ((3-DMAB) 0.93 volt), 4-dimethylaminobenzoic acid ((4-DMAB, 1.07 volts), ethyl p-dimethylaminobenzoate ((EDMAB), 1.07

volts), 2-ethylhexyl p-dimethylaminobenzoate ((EHDMAB), 1.09 volts) and 4-dimethylaminobenzoate ((DMABA), 1.15 volts). The ease of oxidation (and subsequent reactivity) increases as the magnitude of the oxidation decreases. Suitable amine reducing agents in combination with benzoyl peroxide generally include aromatic amines with reduction potentials less than about 1.00 volt vs. 5 SCE. Less effective oxidants than benzoyl peroxide such as lauroyl peroxide (reduction potential = -0.60 volt) are poorer oxidizing agents and subsequently react more slowly with aromatic amine reducing agents. Suitable aromatic amines for lauroyl peroxide will 10 generally include those less than about 0.80 volt vs SCE.

In systems in which two or more reducing agents are present, the relative amounts of each are such that the faster reducing agent(s) is preferably substantially consumed prior to the slower reducing agent(s). Typically, the relative amounts depend on their relative rates of initiation, as well as the types and 15 amounts of other components. Given the disclosure presented herein, one skilled in the art can determine what these amounts should be without undue experimentation. Preferably, the faster reducing agent is present in an amount of about 0.01 wt-% to about 4.0 wt-%, and the slower reducing agent is present in an amount of about 0.01 wt-% to about 4.0 wt-%, based on the total weight of all the 20 components of the dental material. More preferably, the faster reducing agent is present in an amount of about 0.02 wt-% to about 2.0 wt-%, and the slower reducing agent is present in an amount of about 0.02 wt-% to about 2.0 wt-%, based on the total weight of all the components of the dental material. The total amount of reducing agent, based on the weight percent of all the components of the 25 dental material, is preferably within a range of about 0.02 wt-% to about 5.0 wt-%, and more preferably within a range of about 0.1 wt-% to about 3.0 wt-%.

If the reducing agent is a polymerizable tertiary amine, no other reducing agents are required, although others can be used if desired. Surprisingly, a polymerizable tertiary amine reducing agent provides differential initiation rates of 30 the hardening of the resin. The amount of the polymerizable tertiary amine, based on the weight percent of all the components of the dental material, is preferably

within a range of about 1.0 wt-% to about 25.0 wt-%, and more preferably within a range of about 2.0 wt-% to about 15.0 wt-%.

### *Second Initiator System*

5       The dental material of the present invention preferably includes a second initiator system. The second initiator system includes one or more initiators commonly used in free radical polymerization. The function of the second initiator system is to further harden the dental material upon completion of the extended flexible phase work time. The initiators are preferably free radical initiators, which may be activated in a variety of ways, e.g., heat and/or radiation. Thus, the second initiator system can be a redox initiator system, a thermal initiator system (e.g., azo compounds and peroxides), or a photoinitiator system, for example. This final hardening stage of the dental material preferably occurs outside of the oral environment.

15      Preferably, the second initiator system includes one or more photoinitiators. More preferably, the second initiator system includes at least one photoinitiator active in the spectral region of about 300 nanometers (nm) to about 1200 nm and capable of promoting free radical polymerization and/or crosslinking of ethylenically unsaturated moieties upon exposure to light of suitable wavelength and intensity. A wide variety of such photoinitiators can be used. Preferably, they are sufficiently shelf stable and free of undesirable coloration to permit storage and use under typical dental conditions. Visible light photoinitiators are preferred.

20      One type of suitable second initiator system is described in U.S. Pat. No. 5,545,676, which includes a three component or ternary photoinitiator system. This system includes an iodonium salt, e.g., a diaryliodonium salt, which can be a simple salt (e.g., containing an anion such as Cl<sup>-</sup>, Br<sup>-</sup>, I<sup>-</sup>, or C<sub>4</sub>H<sub>9</sub>SO<sub>3</sub><sup>-</sup>) or a metal complex salt (e.g., containing SbF<sub>5</sub>OH<sup>-</sup> or AsF<sub>6</sub><sup>-</sup>). Mixtures of iodonium salts can be used if desired. The second component in this ternary photoinitiator system is a sensitizer, which is capable of light absorption somewhere within the range of wavelengths of about 400 nm to about 1200 nm. The third component in this ternary photoinitiator system is an electron donor and includes amines (including

aminoaldehydes and aminosilanes or other amines as described for the first initiator system), amides (including phosphoramides), ethers (including thioethers), ureas (including thioureas), ferrocene, sulfinic acids and their salts, salts of ferrocyanide, ascorbic acid and its salts, dithiocarbamic acid and its salts, salts of xanthates, salts of ethylene diamine tetraacetic acid and salts of tetraphenylboronic acid.

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Examples of sensitizers suitable for use in the ternary photoinitiator system include ketones, coumarin dyes (e.g., ketocoumarins), xanthene dyes, acridine dyes, thiazole dyes, thiazine dyes, oxazine dyes, azine dyes, aminoketone dyes, porphyrins, aromatic polycyclic hydrocarbons, p-substituted aminostyryl ketone compounds, aminotriaryl methanes, merocyanines, squarylium dyes, and pyridinium dyes. Ketones (e.g., monoketones or alpha-diketones), ketocoumarins, aminoarylketones, and p-substituted aminostyryl ketone compounds are preferred sensitizers. Examples of particularly preferred visible light sensitizers include camphorquinone, glyoxal, biacetyl, 3,3,6,6-tetramethylcyclohexanenedione, 3,3,7,7-tetramethyl-1,2-cycloheptanenedione, 3,3,8,8-tetramethyl-1,2-cyclooctanenedione, 15 3,3,18,18-tetramethyl-1,2-cyclooctadecanenedione, dipivaloyl, benzil, furi, hydroxybenzil, 2,3-butanenedione, 2,3-pentanenedione, 2,3-hexanenedione, 3,4-hexanenedione, 2,3-heptanenedione, 3,4-heptanenedione, 2,3-octanenedione, 4,5-octanenedione, and 1,2-cyclohexanenedione. Of these, camphorquinone is the most preferred sensitizer.

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Yet another type of photoinitiator includes acylphosphine oxides, such as those described in European Pat. Application No. 173567. Suitable acylphosphine oxides are preferably of the general formula  $(R^9)_2-P(=O)-C(=O)-R^{10}$ , wherein each  $R^9$  is individually a hydrocarbon group, preferably an alkyl group, alicyclic group, aryl group, and aralkyl group, any of which can be substituted with a halo-, alkyl- or alkoxy-group, or the two  $R^9$  groups can be joined to form a ring along with the phosphorous atom, and wherein  $R^{10}$  is a hydrocarbon group, preferably, a S-, O-, or N-containing five- or six-membered heterocyclic group, or a  $-Z-C(=O)-P(=O)-(R^9)_2$  group, wherein  $Z$  represents a divalent hydrocarbon group such as alkylene or phenylene having from 2 to 6 carbon atoms. Examples of suitable acylphosphine oxides include bis(2,4,6-trimethylbenzoyl)phenyl phosphine oxide, for example. Optionally, tertiary amine reducing agents may be used in combination with an

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acylphosphine oxide. Illustrative tertiary amines useful in the invention include those described above as well as ethyl 4-(N,N-dimethylamino)benzoate and N,N-dimethylaminoethyl methacrylate.

5 Mono- and di-ketones can also be used as photoinitiators. Examples of such systems are described in U.S. Pat. No. 4,071,424, for example.

Still another class of photoinitiators includes ionic dye-counterion complex initiators that include a borate anion and a complementary cationic dye. Borate anions useful in these photointiators generally can be of the formula  $(R^{11})_4^-$  wherein each  $R^{11}$  is independently an alkyl, aryl, alkaryl, allyl, aralkyl, alkenyl, alkynyl, 10 alicyclic, and saturated or unsaturated heterocyclic groups. Cationic counterions can be cationic dyes, quaternary ammonium groups, transition metal coordination complexes, and the like. Cationic dyes useful as counterions can be cationic methine, polymethine, triarylmethine, indoline, thiazine, xanthene, oxazine or acridine dyes. Quaternary ammonium groups useful as counterions can be trimethylcetylammnonium, cetylpyridinium, and tetramethylammonium. Other 15 organophilic cations can include pyridinium, phosphonium, and sulfonium. Cationic transition metal coordination complexes that may be useful as counterions can be complexes of cobalt, ruthenium, osmium, zinc, iron, and iridium with ligands such as pyridine, 2,2'-bipyridine, 4,4'-dimethyl-2,2'-bipyridine, 1,10- 20 phenanthroline, 3,4,7,8-tetramethylphenanthroline, 2,4,6-tri(2-pyridyl-s-triazine) and related ligands. Borate salt photoinitiators are described, for example, in U.S. Pat. Nos. 4,772,530, 4,954,414, 4,874,450, 5,055,372, and 5,057,393.

Preferred visible light-induced initiators include camphorquinone combined with a suitable hydrogen donor (e.g., an amine such as those described above for the first initiator system), and optionally a diaryliodonium simple or metal complex salts, chromophore-substituted halomethyl-s-triazines, and halomethyl oxadiazoles. Particularly preferred visible light-induced photoinitiators include combinations of an alpha-diketone, e.g., camphorquinone with additional hydrogen donors, and 25 optionally a diaryliodonium salt, e.g., diphenyliodonium chloride, bromide, iodide or hexafluorophosphate.

If present, the initiator is included in the dental material in an amount sufficient to provide the desired rate of hardening. Typically, this amount is

dependent in part on the light source, the thickness of the layer to be exposed to radiant energy, and the extinction coefficient of the initiator. Preferably, an initiator is present at an amount of about 0.01 wt-% to about 5 wt-%, and more preferably from about 0.03 wt-% to about 1.0 wt-%, based on the total weight of  
5 the dental material.

### *Filler System*

A filler system includes one or more fillers and may be selected from one or more materials suitable for incorporation in medical applications, such as the fillers currently used in dental materials. The filler is preferably finely divided with an average particle size (i.e., the longest dimension of the particle, such as the diameter) of no greater than about 10 micrometers and a maximum particle size of no greater than about 50 micrometers. The filler can have a unimodal or polymodal (e.g., bimodal) particle size distribution. The filler can be an inorganic material or it can also be a crosslinked organic material that is insoluble in the resin system. The filler can be radiopaque, radiolucent, or non-radiopaque.  
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Examples of suitable inorganic fillers include naturally occurring or synthetic materials such as quartz, nitrides (e.g., silicon nitride), glasses derived from, for example, Ce, Sb, Sn, Zr, Sr, Ba, or Al, colloidal silica, feldspar, borosilicate glass, kaolin, talc, titania, and zinc glass, zirconia-silica fillers; low Mohs hardness fillers such as those described in U.S. Patent No. 4,695,251; and submicron silica particles (e.g., pyrogenic silicas such as those available under the trade designation "AEROSIL" such as "OX 50," "OX 130," "OX 150," and "OX 200" from Degussa, Ridgefield Park, NJ, and "Cab-O-Sil M5" silica from Cabot Corp., Boston, MA). Examples of suitable organic filler particles include filled or unfilled pulverized polycarbonates, polyepoxides, and the like. Preferred filler particles are quartz, submicron silica, and non-vitreous microparticles of the type described in U.S. Patent No. 4,503,169. Mixtures of these fillers are also contemplated, as well as combination fillers made from organic and inorganic materials. Fluoroaluminosilicate glass fillers, either untreated or silanol treated, are particularly preferred. These glasses have the added benefit of releasing fluoride at the site of dental work when placed in the oral environment.  
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Fillers are preferably chosen to enhance certain properties, such as compressive strength, diametral tensile strength, wear resistance, appearance, translucency, radioopacity, and storage stability of the dental materials, as well as to limit exothermic effects during the initial set phase.

5        Optionally, the surface of the filler particles may be treated with a surface treatment, such as a silane coupling agent, in order to enhance the bond between the filler and the resin system. The coupling agent may be functionalized with reactive curing groups, such as acrylates, methacrylates, and the like.

10      Preferably, the total amount of filler is about 2.0 wt-% to about 80.0 wt-%, based on the total weight of the components of the dental material. More preferably, the total amount of filler is about 30.0 wt-% to about 70.0 wt-%, based on the total weight of the components of the dental material.

#### *Optional Additives*

15      Dental materials of the present invention may additionally include optional adjuvants such as colorants (e.g., pigments conventionally used for shade adjustment), flavorants, medicaments, stabilizers (such as BHT and Tinuvin P), viscosity modifiers, and the like. Such adjuvants may optionally include reactive functionality so that they will be copolymerized with the resin.

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#### *Method of Using the Dental Material*

A preferred method of using the dental material of the invention includes the following steps: 1) make an impression (e.g., an alginate impression) before preparing the tooth (or teeth); 2) prepare the tooth (or teeth); 3) mix the components of the dental material to form a hardenable composition; 4) fill the impression with the hardenable composition (preferably this occurs simultaneously while mixing the components); 5) place the impression over the tooth (or teeth) and allow the hardenable composition to partially harden to a sufficient hardness to be removable from the mouth without being permanently deformed, but not so rigid that the solidified impression locks into undercuts and becomes too hard and brittle to be trimmed.; 6) remove the impression and hardened composition from the mouth; 7) trim the excess hardened material; 8) reseat the restorative in the

mouth to ensure correct fit; 9) allow the restorative to further harden, and if necessary apply heat or light; 10) conduct final trimming, polishing, and cleaning of the restorative; and 11) cement the restorative into place within the mouth.

Typically, the dental material is provided to the dental practitioner in at least two parts, wherein the reducing agent(s) and the oxidizing agent(s), which form the first initiator system are separate. Preferably, there are two parts, each one may include a portion of the resin system and a portion of the filler system. The reducing agent(s) is included in one and the oxidizing agent(s) in the other. The components of the second initiator system can be in either part as long as there is no reaction with the oxidizing and the reducing agents of the first initiator system. Thus, kits are provided by the present invention that include a container for each part. Preferably, two containers are provided in a dual-compartment cartridge, optionally including a static mixing tip, that is used in combination with a hand-held dispenser. Such cartridges and dispensers are commercially available such as that available from Mixpac Systems, Switzerland.

The following examples are given to illustrate, but not limit, the scope of this invention. Unless otherwise indicated, all parts and percentages are by weight and all molecular weights are weight average molecular weight.

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## Test Methods and Preparation of Materials

### Diametral Tensile Strength (DTS) and Compressive Strength (CS) Testing

ADA (“American Dental Association”) specification No. 9 and ADA specification No. 27, respectively, of ISO-test procedure 4049 (1988) were followed for all diametral tensile strength (DTS) and compression strength (CS) testing. Specifically, for CS and DTS testing, the composition was packed into a 4 millimeter (mm) (inside diameter) glass tube, capped with silicone rubber plugs and axially compressed at about 0.28 megapascals (Mpa) for 15 minutes, then light cured for 80 seconds by exposure to two oppositely-disposed VISILUX units (VISILUX 2, 3M St. Paul, MN). Each sample was then irradiated for 90 seconds using a Dentacolor XS unit (Kulzer, Inc., Germany). Cured samples were cut on a diamond saw to form cylindrical plugs 8 mm long for measurement of CS and 2

mm long for measurement of DTS. The plugs were stored in distilled water at 37°C for 24 hours. CS and DTS values for each composition were measured using an INSTRON tester (Instron 4505, Instron Corp., Canton, MA).

5       The Compressive Strength (CS) of these samples was tested on an INSTRON tester with a 10-kN load cell. A total of 5 cylinders of cured composite with about 8 mm length and 4 mm diameter were prepared.

The Diametral Tensile Strength (DTS) of these samples was tested on an INSTRON tester with 10-kN (kilonewton) load cell. A total of 5 cylinders of cured composite with about 2.2 mm length and 4 mm diameter were prepared.

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#### Measurement of Barcol Hardness

Samples of each composite were cured in 2.0 mm thick steel molds sandwiched between polyester (PET) film and glass slides for 60 seconds each with a VISILUX 2 (3M Co., St. Paul, MN). After irradiation, the PET films were 15 removed and the hardness of the sample at the bottom of the mold was measured using a Barber-Coleman Impressor (a hand-held portable hardness tester; Model GYZJ 934-1; Barber-Coleman Co. Industrial Instruments Division, Lovas Park, IN) equipped with an indenter. Top and bottom Barcol hardnesses were measured at 5 minutes and at 24 hours after light exposure.

20       For each composition tested, three readings were taken on the top face and three readings were taken on the bottom face of each sample. The readings were averaged for each composition.

#### Measurement of Shore A Hardness

25       A 10 centimeter (cm) x 10 cm x 1 cm slab of glass was covered with a similar sized stretchable plastic sheet (PARAFILM, American National Can, Neenah, WI). Then a 2.5-mm thick stainless steel ring, having an outer diameter of 57 mm and an inner diameter of 31 mm, was placed on top of the 10 cm x 10 cm slab of glass. A black dual 1:1 compartment cartridge (Part No. CS 050-01-13, 30 Mixpac Systems, Switzerland), containing the restorative paste to be tested, was fitted to a hand held dispenser (Mixpac Systems, Switzerland). The base paste containing a reducing agent was loaded into one of the compartments and the paste

containing an oxidizing agent was loaded into the other compartment prior to testing. The outlet ports of both chambers were drained and then a static mix tip (MBX-4.2-16-S Mixpac Systems, Switzerland) was fitted to the cartridge. This static mixing tip allowed the base paste and the oxidizing agent paste, contained in the two separate chambers of the cartridge, to be mixed homogeneously when the pastes were extruded by means of the hand dispenser.

Approximately 2 mL of paste was extruded inside the inner circle of the previously mentioned stainless steel ring. At this point the stopwatch was started. A second piece of stretchable plastic sheet was placed on top of the fresh paste which had been deposited within the aforementioned stainless steel ring, and the excess paste was squeezed out by means of placing a second 10 cm x 10 cm x 1 cm of glass on top of the stretchable plastic sheet and stainless steel ring. This upper glass slab was removed after 45 seconds. Then a Shore Durometer Hardness Tester, type A2 (The Shore Instrument and Manufacturing Co. in NY, NY) was used to measure hardness as a function of time.

#### Determination of Initial Set Phase Time ( $T_{20}$ )

#### and Extended Flexible Phase Work Time (Delta T)

When time vs. hardness is plotted on an X-Y diagram then an S-shaped curve is obtained. From this chart " $T_{20}$ " and " $T_{80}$ " can be defined as the time required to achieve a Shore A hardness of 20 and 80, respectively. An additional value, "Delta T" is obtained by subtracting  $T_{20}$  from  $T_{80}$ .  $T_{20}$  and Delta T represent the initial set phase time and the extended flexible phase work time, respectively.

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#### Abbreviations/Definitions

BHT	2,6-Di-tert-butyl-4-methylphenol
Bis-GMA	2,2-bis[4(2-hydroxy-3-methacryloyloxy-propyloxy)-phenyl]propane
Bis-EMA-6	6 mole ethoxylated bisphenol A dimethacrylate
BPO	benzoylperoxide
CPQ	camphorquinone
CS	Compressive Strength